



# The Open Cardiovascular Medicine Journal

Content list available at: [www.benthamopen.com/TOCMJ/](http://www.benthamopen.com/TOCMJ/)

DOI: 10.2174/1874192401610010138



## RESEARCH ARTICLE

# The Effects of Levosimendan and Sodium Nitroprusside Combination on Left Ventricular Functions After Surgical Ventricular Reconstruction in Coronary Artery Bypass Grafting Patients

Zeki Temizturk<sup>1</sup>, Davut Azboy<sup>1</sup>, Atakan Atalay<sup>2</sup>, Hakan Atalay<sup>3</sup> and Omer Faruk Dogan<sup>4,\*</sup>

<sup>1</sup>Elazig Education and Training Hospital, Elazig, Turkey

<sup>2</sup>Cukurova University Medical Faculty, Adana, Turkey

<sup>3</sup>Private Mersin Middle East Hospital, Mersin, Turkey

<sup>4</sup>Adana Numune Education and Training Hospital, Adana, Turkey

Received: November 28, 2015

Revised: April 10, 2016

Accepted: April 15, 2015

### Abstract:

#### Objective:

The aim of our study was to research the effects of levosimendan (LS) and sodium nitroprusside (SNP) combination on systolic and diastolic ventricular function after coronary artery bypass grafting (CABG) who required endoventricular patch repair (EVPR).

#### Patients and Methods:

We studied 70 patients with ischemic dilated cardiomyopathy. LS and SNP combination was administered in 35 patients (study group, SG). In the remaining patients, normal saline solution was given (placebo group, PG). Levosimendan (10 $\mu$ gr/kg) started 4 h prior to operation and we stopped LS before the initiation of extracorporeal circulation (ECC). During the rewarming period, we started again levosimendan (10 $\mu$ gr/kg) in combination with SNP (0.1-0.2  $\mu$ gr/kg/min). If mean blood pressure decreased by more than 25% compared with pre-infusion values, for corrected of mean arterial pressure, the volume loading was performed using a 500 ml ringer lactate. Hemodynamic variables, inotrophyc requirement, and laboratory values were recorded.

#### Results:

Five patients died (7.14%) post-surgery (one from SG and 4 from PG) due to low cardiac out-put syndrome (LOS). At the postoperative period, cardiac output and stroke volume index was higher in SG (mean $\pm$ sd;29.1 $\pm$ 6.3 vs. 18.4 $\pm$ 4.9 mL/min<sup>-1</sup>/m<sup>-2</sup> ( $P$ <0.0001)). Stroke volume index (SVI) decreased from 29 $\pm$ 10mL/m<sup>2</sup> preoperatively to 22 $\pm$ 14mL/m<sup>2</sup> in the early postoperative period in group 1. This difference was statistically significant ( $P$ =0.002). Cardiac index was higher in SG (320.7 $\pm$ 37.5 vs. 283.0 $\pm$ 83.9 mL/min<sup>-1</sup>/m<sup>-2</sup> ( $P$ =0.009)). The postoperative inotrophyc requirement was less in SG (5.6 $\pm$ 2.7 vs. 10.4 $\pm$ 2.0 mg/kg,  $P$ < 0.008), and postoperative cardiac enzyme levels were less in SG ( $P$ < 0.01). Ten patients (28.5%) in SG and 21 patients (60%) in PG required inotrophyc support ( $P$ <0.001). We used IABP in eight patients (22.8%) in SG and 17 patients (48.5%) in CG ( $P$ =0.0001).

#### Conclusion:

This study showed that LS and SNP combination impressive increase in left ventricular systolic and diastolic functions including LVEF. The use of this combination achieved more less inotrophics and IABP requirement. We therefore suggest preoperative and peroperative levosimendan and SNP combination.

**Keywords:** Left ventricular functions, Levosimendan, Open heart surgery, Sodium nitroprusside.

\* Address correspondence to this author at the Seyhan Education and Training Hospital, Suleyman Demirel Bulvarı, 01600, Çukurova, Adana, Turkey; Tel: +90 533 481 3056; Fax: +90 322 2143615; E-mail: [ofdogan@hacettepe.edu.tr](mailto:ofdogan@hacettepe.edu.tr)

## INTRODUCTION

Left ventricular aneurysm and cardiomyopathy remains a serious disorder that can lead to intractable congestive heart failure (CHF), and death. The goal of surgical intervention is to correct the size and geometry of the LV, reduce wall tension, and improve ventricular systolic function [1]. EVPR was described as a more physiologic repair than the linear closure technique by Dor and Cooley [2, 3]. Mickleborough and co-workers have introduced a method combining septoplasty and linear resection [4]. Pharmacologic therapies for reducing myocardial afterload and preload has been recommended to induce myocellular and molecular reverse remodeling after surgery [5 - 7]. However, EVPR may be cause of detrimentally affect on LV diastolic properties [8]. Therefore, pharmacologic treatment options are very important after surgery. Levosimendan, as a calcium sensitizer, is used in cardiac surgery to provided the myocardial preconditioning and vasodilating effects without the intracellular calcium accumulation during the ischemia-reperfusion injury. Also, previous investigations have shown that SNP infusion reduces ventricular pre- and afterload after myocardial ischemia by favorably influencing the balance between myocardial oxygen supply and demand [9, 10]. SNP has also suggested for left ventricular relaxation by causing reflex changes in the autonomic nervous system and stimulating catecholamine release. Previous investigation by Freyholdt *et al.* showed that a SNP infusion improved the cardiac index immediately after reperfusion and at the end of surgery [9]. SNP also reduced the cardiac inflammatory response during CABG in patients with severely compromised left ventricular function [10]. According to previous data, SNP may be capable of influencing the myocardial relaxation process under hypoxic conditions and may alter viscoelastic properties of the left ventricle after SVR. To provide detrimental effects of EVPR we hypothesised that pharmacologic preconditioning may be help to increase left ventricular systolic and diastolic functions. To research of the combined drugs effects on left ventricular functions we used LS and SNP for preoperative pharmacologic preconditioning.

## MATERIALS AND METHODS

After the approve of ethics committee, we studied 70 CABG patients with LVA who required EVPR between January 2007 and July 2014. All patients have had previous one or more myocardial infarction. The preoperative patients' characteristics are summarized in Table 1. Criteria for exclusion included a planned concomitant valve procedure, emergency surgery, history of persistent ventricular tachycardia or obstructive cardiomyopathy, myocardial infarction within 72 h before surgery, preoperative inotropic support, or use of an intraaortic balloon pump at the time of surgery. Indications for EVPR were heart failure, angina or a combination of these symptoms. Stroke volume was defined as the difference between end-diastolic volume (EDV) and end-systolic volume (ESV). LV volume measurements were indexed to body surface area and expressed as end-diastolic volume index (EDVI), end-systolic volume index (ESVI), and stroke volume index (SVI). Preoperatively, angiocardiology including left ventricular end-diastolic pressure was calculated.

**Table 1. Baseline characteristics of the study groups.**

Characteristic	LS+SNP (N=50)	CG (N=46)	P Value
Age (years)	59.4±7.1	56.4±5.4	0.095
Sex (males/females)	40/10	35/11	0.228
Smoking	21	20	0.349
Obesity	15	10	0.406
Hypertension	14	6	0.047
Impaired renal function*	4	6	0.94
Diabetes mellitus	8	7	0.90
Mean interval between MI and the operation (years)	3	2.8	0.95
NYHA class II-III	46	43	0.88
Angina pectoris	29	27	0.94
Stable	17	15	-
Unstable	9	7	-
Thromboembolism	0	0	1.0
Mitral regurgitation (grade 1-2)	3	2	0.94
CAD			
2-vessel	16	14	0.94

(Table 1) contd....

Characteristic	LS+SNP (N=50)	CG (N=46)	P Value
3-vessel	34	32	0.92
Mean LVEF	36.7±5.4	38.4±7.2	0.658
Mean LVEDP (mmHg)	16.5±9.7	13.1±5.8	0.546

\*Statistical significant characteristic.

MI: myocardial infarction, LV: left ventricle, NYHA: New York Heart Association, CAD: coronary artery disease, LVEF: left ventricular ejection fraction, LVEDP: left ventricular end diastolic pressure.

In SG, levosimendan (10µg/kg) started 4 h prior to operation. We continued levosimendan administration through central venous catheter until ECC. During the rewarming period, we started levosimendan infusion again (10µgr/kg in combination with SNP (0.1-0.2 µgr/kg/min.)). If mean blood pressure decreased by more than 25% compared with pre-infusion values, the volume loading was performed using a 500 ml ringer lactate for correction of arterial pressure. CG received placebo infusion (0.9% of normal saline) in accordance with LS and SNP infusion protocol. In both groups, echocardiographic examination was done before the discharge from hospital.

### Surgical Procedure

After midline sternal incision, aortic and caval cannulation was performed and CPB was instituted. Cardiac arrest has been provided antegrade and retrograde cold blood cardioplegic solution every 20 min. An aneurysmal sac has been opened and resected in all patients. If there was a thromby, we removed by carefully attention. A pursestring suture was placed around the circumferential scar tissue at the transition zone and a circular Dacron or pericardial patch was secured over the opening after a pursestring suture was tied. If there was a significant septal lesion in addition to the anterior-apex lesions, we used a septal anterior ventricular exclusion procedure after a long left ventriculotomy was made along the left anterior descending artery, an elliptical-shaped Dacron patch was sutured to the transition zone to exclude the akinetic region after broad anteroseptal infarction. Coronary bypass anastomosis were performed using a saphenous vein and a left internal thoracic artery (LITA). During rewarming period, in SG, SNP and levosimendan was administered *via* central venous line at a dose of 0.1-0.2 mg/kg/min., and 5-10µg/kg, respectively. In PG, normal saline solution was given. Proxymal anastomosis were completed using a side clamp. We continued Levosimendan and SNP in ICU for 24 h. The characteristics of surgical procedures are summarized in Table 2. For all patients angiotensin-converting enzyme inhibitor, β-blocker, and diuretics were used. Medication and basic features of the surgical procedures were comparable. There were no differences in heart rate that might have influenced SVI changes after SVR.

**Table 2. Operative data.**

Parameters	LS+SNP(N=50)	CG(N=46)	P value
Mean CPB time (min)	85.1±23.9	94.9±26.2	0.083
Mean aortic cross-clamp time (min)	52.3±14.9	49.7±11.9	0.076
Concomitant CABG	50	46	1.0
Mean number of grafts (grafts per patient)	2.74	3.01	0.670
Urgent operations	8	5	0.802

CPB, cardiopulmonary bypass.

CABG, coronary artery bypass grafting.

### Statistical Analysis

Continuous variables are expressed as mean ± standard deviation. Categorical variables are presented as percentages. The Student *t* test was used to compare preoperative and postoperative continuous variables, the chi-squared test and Fischer's exact test were used for analysis of categorical variables. Univariate regression analysis was used to determine factors associated with early hospital mortality and a low cardiac output state. Survival curves were calculated according to the method of Kaplan - Meier and subgroups were compared using the Log-rank test. Soft ware version 13.0 for Windows was used in data analysis. A significant difference was considered at  $P<0.05$ .

### RESULTS

Five patients died after operations due to LOS (Four in PG (11.4%), and one in SG (2.8%). This was statistically significant ( $P<0.001$ ). The mortality risk factors were duration of ECC (exceeding 220 min. (**P1=0.001**)) and aortic cross clamping (exceeding 100 min.) (**P2=0.001**). Inotrophic support was administered due to LOS in ten patients

(28.5%) from SG, and in 21 patients (60%) in PG, respectively ( $P<0.001$ ). We used IABP in eight patients in SG (22.8%) and in 17 patients (48.5%) in PG ( $P=0.0002$ ). Preoperative lower proportion of New York Heart Association and older age, fibrotic LAD artery, and low EF were the main risk factors for perioperative mortality. LAD revascularization decreased the inotropics and intraaortic balloon pump use both groups ( $P=0.035$ ). But, numbers of coronary artery anastomosis was not statistically significant role of mortality rates ( $P=0.089$ ).

Echocardiographic measurements showed that LVEF increased statistically in all survived patients in SG. However, LVEF did not change in 15 patients (42.8%) in PG. Clinical follow-up demonstrated that the left ventricular EDVI decreased in all survived patients after surgery in both groups.

The LVEF increased from  $34\pm14$  to  $41\pm5$  in SG ( $P=0.001$ ). However, LVEF did not change significantly. The preoperative and postoperative LVEF was  $35\pm16$ ; and  $37\pm3$  in, respectively in PG ( $P>0.05$ ). SVI decreased from  $29\pm10\text{mL/m}^2$  preoperatively to  $22\pm14\text{mL/m}^2$  in the early postoperative period in PG. This was statistically significant ( $P=0.027$ ). In SG patients exhibited an increase in SVI after surgery from  $33\pm14\text{mL/m}^2$  to  $38\pm5\text{mL/m}^2$ . This was statistically significant ( $P<0.001$ ). Table 2 is summarized pre- and postoperative SVI, EDVI, ESVI, LVED and LVESV in both groups in the early postoperative period. The early postoperative data including LOS, mortality rate, intraaortic balloon pump use and arrhythmia have been summarised in Table 3. The changes of left ventricular functions are summarized in Figs. (1 and 2) for SG and PG, respectively.

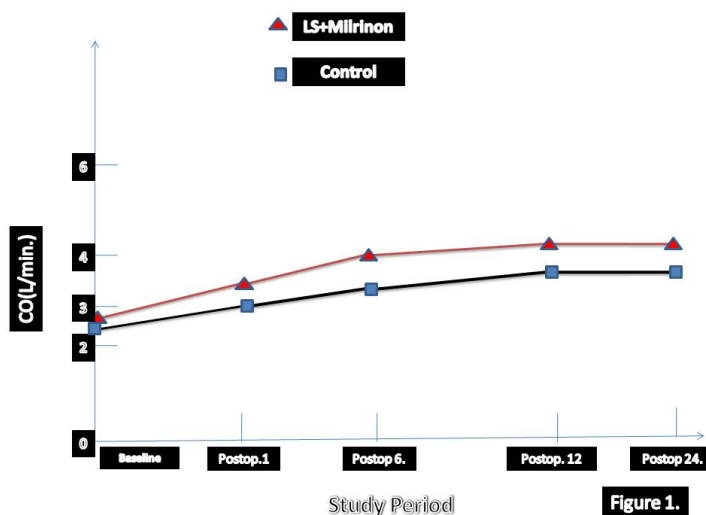
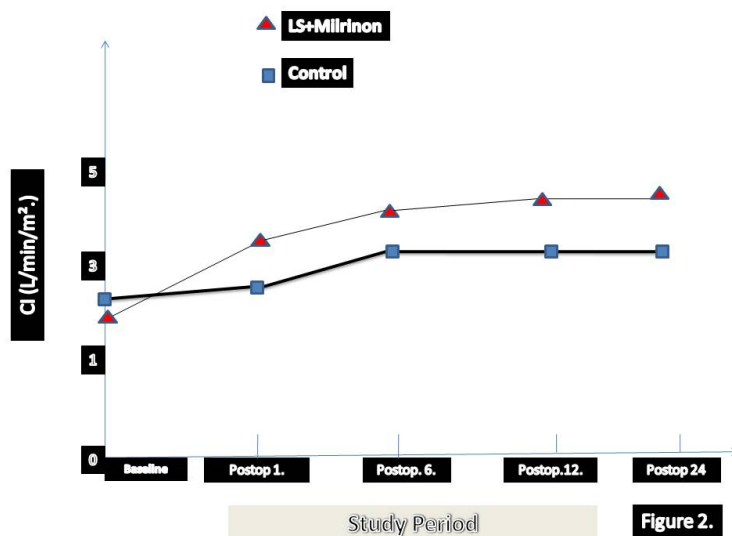


Fig. (1). Shows that the changes of left ventricular functions including postoperative stroke work index, end-systolic volume index, and left ventricular ejection fraction in placebo group. LVEF is stable in 17 patients of placebo group.

Table 3. Early postoperative data.

Parameters	LS+SNP(N=50)	PG (n=46)	P value
Hospital mortality*	2(4%)	4 (8.6%)	0.013
Low cardiac output syndrome*	11 (22%)	23(36.9%)	0.036
Intraaortic balloon pumping*	7 (14%)	16 (34.7%)	0.020
Myocardial infarction	2 (4%)	7 (15.2%)	0.024
Atrial fibrillation*	8 (16%)	17(36.9%)	0.001
Complete AV heart block/	1	2	0.610
Reoperation for bleeding	2	4	0.420
Renal failure*	3 (6%)	12 (27.2%)	0.002
Pneumonia	3 (6%)	3 (6.5%)	0.870

\*Statistical significant charecteristic.



**Fig. (2).** Demonstrates that the changes of left ventricular functions including stroke work index and left ventricular ejection fraction in study group of patients. LVEF is increase significantly after operation in number of patients. LVEF shows an increase from  $36.7\pm 5.4$  to  $43\pm 14\%$ . The mean increment of LVEF is  $18\%\pm 8$  in study group.

Kaplan-Meier Survival Analyses Curve is demonstrated in Fig. (3). During 36 months follow-up, five patients died due to cardiovascular events. Statistical significance has been detected when compared to SVI in the early postoperative period. Post-surgery echocardiographic examinations showed that there was no any statistical difference between the groups ( $P=0.687$ ). The left ventricular end-diastolic and end-systolic dimension were also similar in both groups.

The greater reduction in EDVI and greater increase in EF in early period after surgery have been detected in patients who treated with levosimendan and SNP. According to our results LVEFs were highly correlated to changes in EDVI in SG group. In three times hemodynamic changes has been summarised in Table 4. Univariate and multivariate death from low-output syndrome after surgery has been summarised in Tables 5 and 6, respectively. Long aortic cross-clamp time and concomitant CABG were statistically significant morbidity and mortality factors. We analysed perioperative and postoperative hemodynamic changes over time. Cardiac out-put and CI values were significantly higher in levosimendan group (Table 7).

**Table 4. Hemodynamic changes at three time intervals.**

	SVI decreased	SVI increased	P value
<b>EDVI (mL/m2)</b>			
Pre	111±33	96±25	0.02
Early post	61±10	72±11	0.001
Late post	76±19	88±18.1	0.063
<b>ESVI (mL/m2)</b>			
Pre	56±27	65±19	0.624
Early post	33±8	42±8	0.075
Late post	44±7	47±17	0.080
<b>SVI (mL/m2)</b>	34±7	27±5	0.001
Pre			
Early post	25±6	33±7	0.001
Late post	31±4	34±5	0.675
<b>EF(%)</b>			
Pre	38±4	34±5	0.004
Early post	42±9	40±6	0.624
Late post	43±6	42±8	0.834

SVI : Stroke volume index, EDVI: end-diastolic volume index, ESVI: end-systolic volume index, EF : ejection fraction.

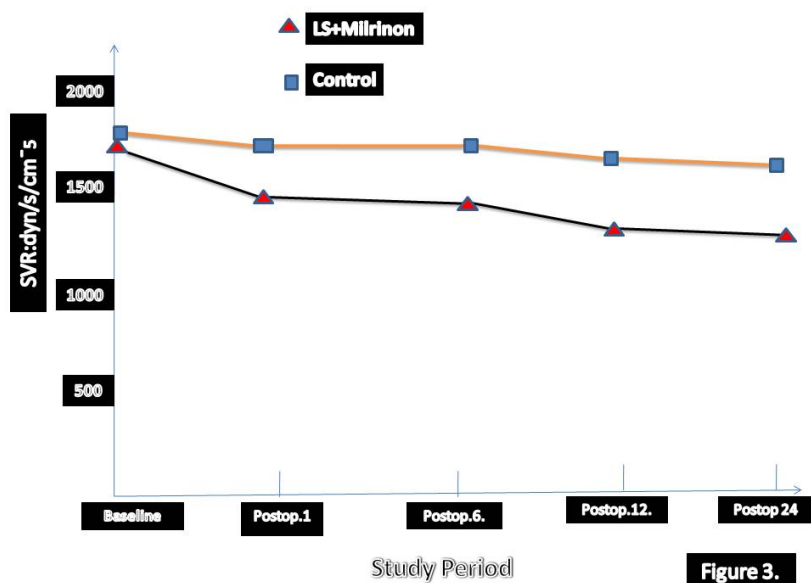


Figure 3.

Fig. (3). Exhibites that Kaplan-Meier survival curve (non-adjusted for demographics) for study and placebo group patients. The mortality rate of placebo group is high during observational period ( $P < 0.01$ ).

Table 5. Univariate analysis for postoperative low cardiac output syndrome and deaths from congestive heart failure.

Variable	LCO (P value)	Deaths from CHF (P value)
<b>Preoperative parameters</b>		
Age 65 years	0.380	0.227
Sex	0.144	1.000
Hypertension	0.470	0.725
Diabetes mellitus	0.469	1.000
Impaired renal function	0.534	0.706
Delay from MI2y.	0.144	0.367
NYHA class	0.190	0.154
Angina pectoris	0.374	0.546
Multivessel CAD	0.512	0.115
LVEDP20 mmHg	0.321	0.015
LVEF <40%	0.580	0.538
<b>Operative parameters</b>		
CPB duration>240 min	0.001	0.002
Aortic cross-clamping 120 min	0.234	0.024
Concomitant CABG	0.037	0.465

Table 6. Multivariate analysis for postoperative low cardiac output syndrome and deaths from congestive heart failure.

Variable	Odds ratio	95%CI	P value
<b>For early LCO</b>			
Left system coronary artery disease	6.9	1.3-19.6	0.024
<b>For death from CHF</b>			
LVEDP >20 mmHg	7.1	1.1-47.6	0.028
<b>CI, confidential interval.</b>			

**Table 7. Peri- postoperative hemodynamic changes over time.**

		8 h After-CPB	12hAfter-CPB	24 h After-CPB	
<b>Heart rate</b>					
Group I	56 ± 14	84 ±6	96 ±11	90 ±9	92 ±7*
Placebo	68 ±18	81±7	93 ±7	97 ± 88	89 ±9*
<b>Mean Arterial Pressure (mmHg)</b>					
Group I	71±8	73± 7	71±7	76 ±13	76 ± 10
Placebo	73±7	72±9	70 ±9	74±9	77 ± 12
<b>Pulmonary Capillary Wedge Pressure*(mmHg)</b>					
Group I	10±2	9±1	8±2	9±1	15 ±2
Placebo	11±3	18±2	17±3	19±2	16 ±2
<b>Central venous pressure (mmHg)</b>					
Group I	12 ±3	13±3	14±3	14±3	13 ±3
Placebo	13±2	14±2	15±3	14±4	14±2
<b>CardiacOutput (L/min)</b>					
Group I	1.8±0.2	2.4 ±0.6*	2.9 ± 0.5*	3.8 ± 1.6*	6.4± 0.5*†
Placebo	1.7 ±0.5	1.9±0.5*	1.9 ±0.4*	2.2 ±0.2*	4.6 ±0.4*
<b>SVR* (dyne/ s/cm-5)</b>					
Group I	1644±166	883±137*	792±152*	788 ±132*	864 ±142*†
Placebo	1672 ± 155	1041± 123*	1040±140*	987±162*	1181 ±182*

Data are mean ± SD. HR, heart rate; MAP, mean arterial pressure; PCWP, pulmonary capillary wedge pressure CVP, central venous pressure; CO: Cardiac output, SVR: systemic vascular resistance.

\*Statistically significant difference compared with base ( $P < 0.05$ ).

### Left Ventricular Size and Functions in Long-Term Follow-Up

In survived patients, echocardiograms were available in the first week post-EVPR, and a median of 6, 4 months (range 3-25 months). Five patients died because of cardiovascular events during the follow-up period (three patients from PG and, two patients from SG). The remaining 60 patients did not differ in clinical and hemodynamic profiles when we compared echocardiographic controls. Patients in whom SV decreased initially had larger SV at baseline compared with those in whom SV increased initially; they also had larger EDVI and ESVI and greater EF. According to Marui *et al.*, our research showed that the SVI reduction was temporary, and patients who have lower SV showed recovery during the follow-up, so that at the time of chronic follow-up there was no significant difference in SV between the two groups [11, 12]. However, the reduction in EDVI and ESVI persisted along time in the PG. These findings indicate that different baseline loading conditions may affect the response of cardiac function to SVR in SG.

### DISCUSSION

Postoperative ventricular failure is a life threatening problem in CABG patients. Impairment of ventricular function produces a reduction in peripheral oxygen transport, which, in turn, leads to a prolonged stay in the ICU. Myocardial  $\beta$ -adrenergic receptor desensitization occurs chronically in patients with congestive heart failure and acutely after ECC, thereby limiting the efficacy of  $\beta$ -adrenergic stimulants for post-bypass cardiac failure. We presented the effects of levosimendan and sodium nitroprusside combination in CABG patients who required EVPR. Our research demonstrated that older age, a low LVEF as independent risk factors. Also, our study was clearly showed that NYHA class III and IV status, longer aortic cross clamp and total ECC time were the most important factors of early mortality in CABG patients who underwent EVPR. To the best of our knowledge, for the first time, we investigated the effects of the use of levosimendan and SNP combination on left ventricular systolic and diastolic functions including SVI, EDVI, ESVI and LVEF in our CABG patients who have LF aneurysm requiring surgical ventricular reconstruction.

Levosimendan is a calcium sensitizers, that increasemyocardial contractility without increasing intracellular calcium, has emerged as an inotropic agent. LS has been used for myocardial preconditioning without the intracellular calcium accumulation during the ischemia-reperfusion injury [13]. Because of adenosine triphosphate (ATP) production, LS provides coronary artery dilationand myocyte mitochondrial activation. These beneficial effects work synergistically with calcium sensitization to ventricular myocardial improvement [13]. Tritapepe *et al.* reported that levosimendan pre-treatment improves outcomes in patients undergoing CABG patients [14]. Ikonomidis et colleagues showed that treatment with LS improved coronary artery flow and microcirculation in parallel with an improvement in

cardiac performance and neuro-hormonal activation in patients with advanced heart failure [15]. Levin's and Malliotakis's study demonstrated that levosimendan decreased the mortality in high risk patients undergoing CABG with impaired left ventricular function [16, 17]. Our previously published study exhibited that levosimendan decreased atrial fibrillation after CABG [18].

Sodium nitroprusside could potentially influence the left ventricular relaxation by several mechanism in patients with myocardial ischemia [19].

We hypothesised that after EVPR the left ventricular volume may be decreased and diastolic filling rates fell significantly during LV diastole, and did not change and fell significantly, during the last third of diastole. Therefore, these changes in filling rates may contribute to the shifts in the diastolic pressure-volume relation seen in our patients when we administer SNP and levosimendan for ventricular preconditioning. Our study showed that the ventricular diastolic filling rates increased slightly with our pharmacologic approaches in CABG patients who underwent EVPR. The results of our study showed that EF and resting SV was more frequently increased after SVR in SG. In fact, in data of PG cohort showed an average 8 mL/m<sup>2</sup> (approximately 14%) decrease in resting SV.

Clinical studies have showed EVPR frequently decreased EDV [17 - 20]. Vasodilating effects of SNP and its positive impact on afterload after cross clamping increased the LVEF, EDV, and ESV in SG. Our results demonstrated preoperative levosimendan preconditioning and SNP use directly affected on index of LV pump function.

Our study demonstrated that administration of LS combined with SNP increased ventricular relaxation and LVEF after EVPR in dilated ischemic cardiomyopathy. LS and SNP decreased intraaortic balloon pump and inotropic agent requirement. LS and SNP may influence the left ventricular systolic and diastolic pressure-volume relationships. Likewise, the pressure and blood flow in the coronary arteries and their intramural branches could affect the compliance of the left ventricular wall and influence the pressure-volume curve on this basis. Overall, our estimates of mean postoperative SVR changes in SV reported in the literature range from an increase of 15 mL to decreases of 15 mL or greater with, as noted above, a majority of studies showing reductions [21, 22].

### Study Limitations

Our data set concerning the impact of EVPR on left ventricular size and function was in the early postoperative period, with a smaller database. Comparison of resting LV size and function between preoperative and early postoperative conditions provides important information. The previous study confirmed that important specific factor was defined as ventricular filling characteristics and the LV systolic and diastolic properties. The left ventricular clarification such as ventricular diastolic properties associated with better hemodynamic effects of SVR and links between hemodynamic effects and clinical outcomes could help define appropriate patient selection criteria for these pharmacologic approaches and surgical procedures. Despite our limitations, the current results advocate routine use of levosimendan protocol in patients with low LVEF undergoing OPCABG surgery.

### CONCLUSION

Preoperative levosimendan and peroperative SNP combined with levosimendan administration in CABG patients with low LVEF undergoing ECC was associated with greater improvement in hemodynamic stability after surgery and early postoperative period. It shows faster onset of action, lower use of inotropic support, and IABP use including shorter ICU and hospital stay and lower rate of complications as well as morbidity. The present study analyzed the early changes in SVI after SVR and attempted to understand the apparent paradox of improved volumes reduction and LVEF with the frequently observed reduction in stroke volume after SVR.

### CONFLICT OF INTEREST

The authors confirm that this article content has no conflict of interest.

### ACKNOWLEDGEMENTS

Declared none.

### REFERENCES

- [1] Cooley DA, Collins HA, Morris GC Jr, Chapman DW. Ventricular aneurysm after myocardial infarction; surgical excision with use of temporary cardiopulmonary bypass. *J Am Med Assoc* 1958; 167(5): 557-60. [<http://dx.doi.org/10.1001/jama.1958.02990220027008>] [PMID: 13538738]



- [2] Dor V, Saab M, Coste P, Kornaszewska M, Montiglio F. Left ventricular aneurysm: a new surgical approach. *Thorac Cardiovasc Surg* 1989; 37(1): 11-9. [http://dx.doi.org/10.1055/s-2007-1013899] [PMID: 2522252]
- [3] Cooley DA. Ventricular endoaneurysmorrhaphy: a simplified repair for extensive postinfarction aneurysm. *J Card Surg* 1989; 4(3): 200-5. [http://dx.doi.org/10.1111/j.1540-8191.1989.tb00282.x] [PMID: 2535278]
- [4] Mickleborough LL, Carson S, Ivanov J. Repair of dyskinetic or akinetic left ventricular aneurysm: results obtained with a modified linear closure. *J Thorac Cardiovasc Surg* 2001; 121(4): 675-82. [http://dx.doi.org/10.1067/mtc.2001.112633] [PMID: 11279407]
- [5] Mickleborough LL, Maruyama H, Liu P, Mohamed S. Results of left ventricular aneurysmectomy with a tailored scar excision and primary closure technique. *J Thorac Cardiovasc Surg* 1994; 107(3): 690-8. [PMID: 8127098]
- [6] Liden H, Karason K, Bergh CH, Nilsson F, Koul B, Wiklund L. The feasibility of left ventricular mechanical support as a bridge to cardiac recovery. *Eur J Heart Fail* 2007; 9(5): 525-30. [http://dx.doi.org/10.1016/j.ejheart.2006.12.003] [PMID: 17257894]
- [7] Jones RH, White H, Velazquez EJ, et al. STICH (Surgical Treatment for Ischemic Heart Failure) trial enrollment. *J Am Coll Cardiol* 2010; 56(6): 490-8. [http://dx.doi.org/10.1016/j.jacc.2009.11.102] [PMID: 20670760]
- [8] Marui A, Nishina T, Saji Y, et al. Significance of left ventricular diastolic function on outcomes after surgical ventricular restoration. *Ann Thorac Surg* 2010; 89(5): 1524-31. [http://dx.doi.org/10.1016/j.athoracsur.2010.01.067] [PMID: 20417772]
- [9] Freyholdt T, Massoudy P, Zahler S, et al. Beneficial effect of sodium nitroprusside after coronary artery bypass surgery: pump function correlates inversely with cardiac release of proinflammatory cytokines. *J Cardiovasc Pharmacol* 2003; 42(3): 372-8. [http://dx.doi.org/10.1097/00005344-200309000-00008] [PMID: 12960682]
- [10] Massoudy P, Zahler S, Freyholdt T, et al. Sodium nitroprusside in patients with compromised left ventricular function undergoing coronary bypass: reduction of cardiac proinflammatory substances. *J Thorac Cardiovasc Surg* 2000; 119(3): 566-74. [http://dx.doi.org/10.1016/S0022-5223(00)70138-3] [PMID: 10694618]
- [11] Marui A, Nishina T, Saji Y, et al. Significance of left ventricular diastolic function on outcomes after surgical ventricular restoration. *Ann Thorac Surg* 2010; 89(5): 1524-31. [http://dx.doi.org/10.1016/j.athoracsur.2010.01.067] [PMID: 20417772]
- [12] Leslie J, Brister N, Levy JH, et al. Treatment of postoperative hypertension after coronary artery bypass surgery. Double-blind comparison of intravenous isradipine and sodium nitroprusside *Circulation* 1994; 90(5): 256-61.
- [13] Deakin CD, Petley GW, Smith D. Pharmacological vasodilatation improves efficiency of rewarming from hypothermic cardiopulmonary bypass. *Br J Anaesth* 1998; 81(2): 147-51. [http://dx.doi.org/10.1093/bja/81.2.147] [PMID: 9813513]
- [14] Tritapepe L, De Santis V, Vitale D, et al. Levosimendan pre-treatment improves outcomes in patients undergoing coronary artery bypass graft surgery. *Br J Anaesth* 2009; 102(2): 198-204. [http://dx.doi.org/10.1093/bja/aen367] [PMID: 19151048]
- [15] Ikonomidis I, Parissis JT, Paraskevaidis I, et al. Effects of levosimendan on coronary artery flow and cardiac performance in patients with advanced heart failure. *Eur J Heart Fail* 2007; 9(12): 1172-7. [http://dx.doi.org/10.1016/j.ejheart.2007.10.002] [PMID: 18062901]
- [16] Levin R, Degrange M, Del Mazo C, Tanus E, Porcile R. Preoperative levosimendan decreases mortality and the development of low cardiac output in high-risk patients with severe left ventricular dysfunction undergoing coronary artery bypass grafting with cardiopulmonary bypass. *Exp Clin Cardiol* 2012; 17(3): 125-30. [PMID: 23620700]
- [17] Malliotakis P, Xenikakis T, Linardakis M, Hassoulas J. Haemodynamic effects of levosimendan for low cardiac output after cardiac surgery: a case series. *Hellenic J Cardiol* 2007; 48(2): 80-8. [PMID: 17489345]
- [18] Dogan OF. Levosimendan decreased atrial fibrillation in our CABG patients with poor left ventricle. *Heart Surg Forum* 2013; 16(5): 287-94.
- [19] Di Donato M, Fantini F, Toso A, et al. Impact of surgical ventricular reconstruction on stroke volume in patients with ischemic cardiomyopathy. *J Thorac Cardiovasc Surg* 2010; 140(6): 1325-31.e1, 2. [http://dx.doi.org/10.1016/j.jtcvs.2010.01.031] [PMID: 20381078]
- [20] Burkhoff D, Wechsler AS. Surgical ventricular remodeling: a balancing act on systolic and diastolic properties. *J Thorac Cardiovasc Surg* 2006; 132(3): 459-63. [http://dx.doi.org/10.1016/j.jtcvs.2006.04.045] [PMID: 16935094]
- [21] Artrip JH, Oz MC, Burkhoff D. Left ventricular volume reduction surgery for heart failure: a physiologic perspective. *J Thorac Cardiovasc Surg* 2001; 122(4): 775-82. [http://dx.doi.org/10.1067/mtc.2001.116208] [PMID: 11581612]

- [22] Ratcliffe MB, Guy TS. The effect of preoperative diastolic dysfunction on outcome after surgical ventricular remodeling. *J Thorac Cardiovasc Surg* 2007; 134(2): 280-3.  
[<http://dx.doi.org/10.1016/j.jtcvs.2007.01.088>] [PMID: 17662762]

---

© Temizturk *et al.*; Licensee *Bentham Open*.

This is an open access article licensed under the terms of the Creative Commons Attribution-Non-Commercial 4.0 International Public License (CC BY-NC 4.0) (<https://creativecommons.org/licenses/by-nc/4.0/legalcode>), which permits unrestricted, non-commercial use, distribution and reproduction in any medium, provided the work is properly cited.